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### Status: Path 1 of [Dialog Information Services via Modem]
### Status: Initializing TCP/IP using (UseTelnetProto 1 ServiceID pto-dialog)
Trying 31060000009999...Open
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PLEASE LOGON:
 ****** HHHHHHHH SSSSSSSS?
### Status: Signing onto Dialog
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ENTER PASSWORD:
 ****** HHHHHHHH SSSSSSS? *******
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### Status: Connected
Dialog level 02.03.27D
Last logoff: 15mar02 07:52:56
Logon file405 16apr02 12:42:18
           *** ANNOUNCEMENT ***
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--U.S. Patents Fulltext (File 654) has been redesigned with
new search and display features. See HELP NEWS 654 for
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--Dialog NewsRoom is now available. BEGIN NEWSROOM
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(below) for individual file numbers.
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--CLAIMS/US Patents (Files 340,341, 942) have been enhanced
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-- Important news for public and academic
libraries. See HELP LIBRARY for more information.
-- Important Notice to Freelance Authors--
See HELP FREELANCE for more information
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***Dialog NewsRoom - 2001 Archive (File 994)
***Dialog NewsRoom - 2000 Archive (File 995)
***AGROProjects (File 235)
***TRADEMARKSCAN-Finland (File 679)
***TRADEMARKSCAN-Japan (File 669)
***TRADEMARKSCAN-Norway (File 678)
***TRADEMARKSCAN-Sweden (File 675)
UPDATING RESUMED
***Delphes European Business (File 481)
RELOADED
***U.S. Patents Fulltext 1976-current (File 654)
***Population Demographics (File 581)
***CLAIMS/US PATENTS (Files 340, 341, 942)
***Kompass Western Europe (590)
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A. 1

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***D&B - Dun's Market Identifiers (516)
REMOVED
***U.S. Patents Fulltext 1980-1989 (File 653)
***Washington Post (File 146)
***Books in Print (File 470)
***Court Filings (File 793)
***Microcomputer Software Guide Online (File 278)
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   broad spectrum of news wires.
     >>> Enter BEGIN HOMEBASE for Dialog Announcements <<<
           of new databases, price changes, etc.
KWIC is set to 50.
HILIGHT set on as '*'
PICKS is set ON as an alias for 5,55,159,143,358,340,344,348,351,352,447,72,73,154,155,34
9.
SYSTEM: HOME
Cost is in DialUnits
Menu System II: D2 version 1.7.8 term=ASCII
                     *** DIALOG HOMEBASE(SM) Main Menu ***
 Information:
  1. Announcements (new files, reloads, etc.)
  2. Database, Rates, & Command Descriptions
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                           /L = Logoff
Enter an option number to view information or to connect to an online
 service. Enter a BEGIN command plus a file number to search a database
(e.g., B1 for ERIC).
?b pick
>>>"PICK" is not a valid category or service name
>>>No valid files specified
?b picks
            351 is unauthorized
>>>
            352 is unauthorized
>>>2 of the specified files are not available
       16apr02 12:42:29 User243038 Session D96.1
                    0.309 DialUnits FileHomeBase
            $0.00
     $0.00 Estimated cost FileHomeBase
     $0.04 TELNET
     $0.04 Estimated cost this search
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SYSTEM:OS - DIALOG OneSearch
                                1969-2002/Apr W1
  File
         5:Biosis Previews(R)
         (c) 2002 BIOSIS
                               1993-2002/Apr W1
  File
        55:Biosis Previews(R)
         (c) 2002 BIOSIS
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*File 159: For UDs information please see Help News159.
  File 143:Biol. & Agric. Index 1983-2002/Mar
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  File 358:Current BioTech Abs 1983-2001/Oct
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  File 340:CLAIMS(R)/US Patent 1950-02/Apr 04
         (c) 2002 IFI/CLAIMS(R)
*File 340: Both the application and grant publication levels for a
patent are in a single record. See HELP NEWS 340 for details. File 344:CHINESE PATENTS ABS APR 1985-2002/MAR
         (c) 2002 EUROPEAN PATENT OFFICE
  File 348:EUROPEAN PATENTS 1978-2002/APR W01
         (c) 2002 European Patent Office
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see Help News73.
                      1990-2002/Apr W1
  File 154:MEDLINE(R)
  File 155:MEDLINE(R) 1966-2002/Apr W1
  File 349:PCT FULLTEXT 1983-2002/UB=20020411,UT=20020404
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?s collagen
      S1 390127 COLLAGEN
?s collagen?
      S2 462795 COLLAGEN?
?s type I and s2
                 TYPE I
            1534
          462795 S2
             262 TYPE I AND S2
      S3
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           97768 ANGIOGENESIS
              40 S4 AND ANGIOGENESIS
      S6
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Processing
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Completed processing all files
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                  S6
         4897395
                  INHIBIT?
                  S6 AND INHIBIT?
      $7
              10
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              10
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2551850 ANTIBOD?

DIALOG(R) File 5:Biosis Previews(R) (c) 2002 BIOSIS. All rts. reserv.

10530456 BIOSIS NO.: 199699151601

Involvement of the transcription factor NF-kappa-B in tubular morphogenesis of human microvascular endothelial cells by oxidative stress.

AUTHOR: Shono Tadahisa; Ono Mayumi; Izumi Hiroto; Jimi Sei-Ichiro; Matsushima Kouji; Okamoto Takashi; Kohno Kimitoshi; Kuwano Michihiko(a) AUTHOR ADDRESS: (a) Dep. Biochemistry, Kyushu Univ. Sch. Med., Maidashi, Fukuoka 812-82**Japan

JOURNAL: Molecular and Cellular Biology 16 (8):p4231-4239 1996

ISSN: 0270-7306

DOCUMENT TYPE: Article RECORD TYPE: Abstract LANGUAGE: English

ABSTRACT: Oxygen radicals are induced under various pathologic conditions associated with neovascularization. Oxygen radicals modulate *angiogenesis* in cultured human microvascular endothelial cells by an unknown mechanism. Treatment of human microvascular endothelial cells for 15 min with 0.1 to 0.5 mM hydrogen peroxide (H-20-2) or 100 U of tumor necrosis factor alpha per ml induced tubular morphogenesis in type I collagen gels. Gel shift assays with nuclear extracts demonstrated that H-20-2 increases the binding activities of two transcription factors, NF-kappa-B and AP-1, but not of Sp1. Tumor necrosis factor alpha increased the binding activities of all three factors. A supershift assay with specific *antibodies* against JunB, JunD, and c-jun (Jun family) showed that the *antibody* against c-jun supershifted the AP-1 complex after H-20-2 treatment. Coadministration of the antisense sequence of NF-KB *inhibited* H-20-2-dependent tubular morphogenesis, and the antisense c-Jun oligonucleotide caused partial *inhibition*. The angiogenic factor responsible for H-20-2-induced tubular morphogenesis was examined. Cellular mRNA levels of vascular endothelial growth factor and interleukin-8 (IL-8), but not those of transforming growth factor et, were increased after treatment with 0.5 mM H-20-2. Coadministration of anti-IL-8 *antibody* *inhibited* tubular morphogenesis enhanced by H-20-2, and IL-8 itself also enhanced the formation of tube-like structures. Treatment with antisense NF-KB oligonucleotide completely blocked H-20-2-dependent IL-8 production by endothelial cells. The tubular morphogenesis of vascular endothelial cells after treatment with oxidative stimuli and its possible association with NF-kappa-B and IL-8, is examined.

REGISTRY NUMBERS: 7722-84-1: HYDROGEN PEROXIDE DESCRIPTORS:

MAJOR CONCEPTS: Biochemistry and Molecular Biophysics; Blood and Lymphatics (Transport and Circulation); Cardiovascular System (Transport and Circulation); Development; Endocrine System (Chemical Coordination and Homeostasis); Genetics; Molecular Genetics (Biochemistry and Molecular Biophysics)

BIOSYSTEMATIC NAMES: Hominidae--Primates, Mammalia, Vertebrata, Chordata, Animalia

ORGANISMS: Hominidae (Hominidae)

BIOSYSTEMATIC CLASSIFICATION (SUPER TAXA): animals; chordates; humans; mammals; primates; vertebrates

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HYDROGEN PEROXIDE
  CHEMICALS & BIOCHEMICALS:
                       HYDROGEN PEROXIDE; INTERLEUKIN-8; MESSENGER RNA;
 MISCELLANEOUS TERMS:
   NEOVASCULARIZATION; TUMOR NECROSIS FACTOR-ALPHA; *TYPE I COLLAGEN*;
    VASCULAR ENDOTHELIAL GROWTH FACTOR
CONCEPT CODES:
          Genetics and Cytogenetics-Human
  03508
         Biochemistry-Gases (1970-)
  10012
         Replication, Transcription, Translation
  10300
         Cardiovascular System-Physiology and Biochemistry
  14504
         Blood, Blood-Forming Organs and Body Fluids-Lymphatic Tissue and
  15008
             Reticuloendothelial System
  17002
          Endocrine System-General
          Developmental Biology-Embryology-Morphogenesis, General
  25508
         Cytology and Cytochemistry-Human
  02508
         Biochemical Studies-General
  10060
         Biochemical Studies-Nucleic Acids, Purines and Pyrimidines
  10062
         Biochemical Studies-Proteins, Peptides and Amino Acids
  10064
         Biophysics-Molecular Properties and Macromolecules
  10506
BIOSYSTEMATIC CODES:
  86215 Hominidae
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S1
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S2
S3
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               TYPE I AND S2
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               TYPE I COLLAGEN
S4
S5
          Ω
               S4 AND INHIBIT? ANGIOGENESIS
          40 S4 AND ANGIOGENESIS
S6
S7
          10 S6 AND INHIBIT?
          2 S7 AND ANTIBOD?
S8
              RD (unique items)
           1
S9
?t s7/5/all
           (Item 1 from file: 5)
 7/5/1
DIALOG(R) File 5:Biosis Previews(R)
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          BIOSIS NO.: 200000012661
12259159
*Inhibition* by vasoactive intestinal polypeptide (VIP) of *angiogenesis*
  induced by murine Colon 26-L5 carcinoma cells metastasized in liver.
AUTHOR: Ogasawara Masaru; Murata Jun(a); Kamitani Yukio; Hayashi Kazuko;
  Saiki Ikuo
AUTHOR ADDRESS: (a) Department of Pathogenic Biochemistry, Institute of
  Natural Medicine, Toyama Medical and Pharmaceutical University, 2630
  Sugitani, Toyama, 930-0194**Japan
JOURNAL: Clinical & Experimental Metastasis 17 (4):p283-291 June, 1999
ISSN: 0262-0898
DOCUMENT TYPE: Article
RECORD TYPE: Abstract
LANGUAGE: English
SUMMARY LANGUAGE: English
ABSTRACT: We investigated the effect of VIP on the liver metastases and
  *angiogenesis* by Colon 26-L5 carcinoma cells in mice. Daily systemic
  administration of VIP, beginning 3 days after tumor inoculation into a
  portal vein of mice, *inhibited* significantly the development of their
  liver metastases. Immunohistochemical staining for factor VIII-related
  antigen in the sections of liver metastases showed that the systemic
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angiogenesis by Colon 26-L5 carcinoma cells in mice. Daily systemic administration of VIP, beginning 3 days after tumor inoculation into a portal vein of mice, *inhibited* significantly the development of their liver metastases. Immunohistochemical staining for factor VIII-related antigen in the sections of liver metastases showed that the systemic administration of VIP caused significant prevention of *angiogenesis* within tumor masses. VIP (10-10 to 10-6 M) *inhibited* the invasion of reconstituted basement membrane (Matrigel) by hepatic sinusoidal endothelial (HSE) cells in a concentration-dependent manner in a Transwell chamber assay in vitro and achieved approximately 50% reduction of control at 10-6 M. VIP (10-6 M) also significantly suppressed the haptotactic migration of HSE cells to fibronectin, laminin or type I collagen substrates with a similar *inhibition* rate to the invasion assay. Exposure of VIP to HSE cell s induced accumulation of intracellular cAMP in a concentration-dependent manner. The *inhibitory*

effect of VIP (10-6 M) on HSE cell migration was significantly abrogated in the presence of 3 X 10-6 M H-89, a cAMP-dependent protein kinase *inhibitor*. VIP (10-6 M) *inhibited* the morphogenesis of HSE cells into capillary-like structures on Matrigel-coated wells. VIP did not affect the proliferation of HSE cells and the production of gelatinases in HSE cells in vitro at the concentrations used in the invasion assay. These observations suggest that the anti-metastatic effect of VIP on liver metastases by Colon 26-L5 carcinoma cells in mice is partly due to the prevention of tumor *angiogenesis* probably through suppression of the motility of endothelial cells.

REGISTRY NUMBERS: 60-92-4: CYCLIC AMP; 9040-48-6: GELATINASE; 37221-79-7: VASOACTIVE INTESTINAL POLYPEPTIDE DESCRIPTORS: MAJOR CONCEPTS: Digestive System (Ingestion and Assimilation); Tumor BIOSYSTEMATIC NAMES: Muridae--Rodentia, Mammalia, Vertebrata, Chordata, Animalia ORGANISMS: Balb/c mouse (Muridae) -- animal model; colon 26-L5 cell line (Muridae) -- carcinoma cells ORGANISMS: PARTS ETC: hepatic sinusoidal endothelial cells--digestive system, haptotactic migration, morphogenesis, proliferation, tube formation BIOSYSTEMATIC CLASSIFICATION (SUPER TAXA): Animals; Chordates; Mammals; Nonhuman Mammals; Nonhuman Vertebrates; Rodents; Vertebrates DISEASES: colon cancer--digestive system disease, neoplastic disease; liver metastasis--digestive system disease, histopathology, neoplastic disease CHEMICALS & BIOCHEMICALS: cAMP {cyclic AMP}--concentration-dependent intracellular accumulation; fibronectin; gelatinase; laminin; *type I collagen*; vasoactive intestinal polypeptide--anti-angiogenic effect , anti-metastatic effect METHODS & EQUIPMENT: immunohistochemistry--histochemical method tumor *angiogenesis*--*inhibition* MISCELLANEOUS TERMS: ALTERNATE INDEXING: Colonic Neoplasms (MeSH); Liver Neoplasms (MeSH) CONCEPT CODES: Neoplasms and Neoplastic Agents-General 24002 Digestive System-General; Methods 14001 BIOSYSTEMATIC CODES:

7/5/2 (Item 2 from file: 5)
DIALOG(R)File 5:Biosis Previews(R)
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Muridae

11974220 BIOSIS NO.: 199900227533

Signaling via fibroblast growth factor receptor-1 is dependent on extracellular matrix in capillary endothelial cell differentiation.

AUTHOR: Kanda Shigeru; Tomasini-Johansson Bianca; Klint Peter; Dixelius Johan; Rubin Kristofer; Claesson-Welsh Lena(a)

AUTHOR ADDRESS: (a) Department of Medical Biochemistry and Microbiology, Biomedical Center, S-751 23, Uppsala**Sweden

JOURNAL: Experimental Cell Research 248 (1):p203-213 April 10, 1999

ISSN: 0014-4827

86375

DOCUMENT TYPE: Article RECORD TYPE: Abstract LANGUAGE: English

SUMMARY LANGUAGE: English

ABSTRACT: Differentiation of endothelial cells, i.e., formation of a vessel lumen, is a prerequisite for *angiogenesis*. The underlying molecular mechanisms are ill defined. We have studied a brain capillary endothelial cell line (IBEC) established from H-2Kb-tsA58 transgenic mice. These cells form hollow tubes in three-dimensional type I collagen gels in response to fibroblast growth factor-2 (FGF-2). Culture of IBEC on collagen gels in the presence of FGF-2 protected cells from apoptosis and allowed tube formation (i.e., differentiation) but not growth of the cells. FGF-induced differentiation, but not cell survival, was

inhibited by treatment of the cells with an anti-betal-integrin IgG. Changes in integrin expression in the collagen-gel cultures could not be detected. Rather, cell-matrix interactions critical for endothelial cell differentiation were created during the culture, as indicated by the gradual increase in tyrosine phosphorylation of focal adhesion kinase in the collagen-gel cultures. Inclusion of laminin in the collagen gels led to FGF-2-independent formation of tube structures, but cells were not protected from apoptosis. These data indicate that FGF receptor-1 signal transduction in this cell model results in cell survival. Through mechanisms dependent on cell-matrix interactions, possibly involving the alpha3beta1-integrin and laminin produced by the collagen-cultured IBE cells, FGF stimulation also leads to differentiation of the cells.

REGISTRY NUMBERS: 60-18-4Q: TYROSINE; 556-03-6Q: TYROSINE; 9031-44-1: KINASE DESCRIPTORS: MAJOR CONCEPTS: Cardiovascular System (Transport and Circulation); Cell BIOSYSTEMATIC NAMES: Muridae -- Rodentia, Mammalia, Vertebrata, Chordata, Animalia ORGANISMS: IBEC cell line (Muridae) -- brain capillary endothelial cell ORGANISMS: PARTS ETC: capillary endothelial cell--circulatory system, differentiation BIOSYSTEMATIC CLASSIFICATION (SUPER TAXA): Animals; Chordates; Mammals; Nonhuman Mammals; Nonhuman Vertebrates; Rodents; Vertebrates CHEMICALS & BIOCHEMICALS: extracellular matrix; fibroblast growth factor receptor-1; fibroblast growth factor-2; focal adhesion kinase --tyrosine phosphorylation; *type I collagen*; tyrosine MISCELLANEOUS TERMS: *angiogenesis*; signal transduction CONCEPT CODES: 02506 Cytology and Cytochemistry-Animal Biochemical Studies-General 10060 Enzymes-General and Comparative Studies; Coenzymes 10802 Cardiovascular System-General; Methods 14501 Endocrine System-General 17002 Nervous System-General; Methods 20501 BIOSYSTEMATIC CODES: Muridae 86375 (Item 3 from file: 5) DIALOG(R)File 5:Biosis Previews(R) (c) 2002 BIOSIS. All rts. reserv. BIOSIS NO.: 199800267849 *Inhibition* of *angiogenesis* on glycated collagen lattices. AUTHOR: Kuzuya M(a); Satake S; Ai S; Asai T; Kanda S; Ramos M A; Miura H; Ueda M; Iquchi A AUTHOR ADDRESS: (a) Dep. Geriatr., Nagoya Univ. Sch. Med., 65 Tsuruma-cho, Showa-ku, Nagoya 466-8550**Japan JOURNAL: Diabetologia 41 (5):p491-499 May, 1998 ISSN: 0012-186X DOCUMENT TYPE: Article RECORD TYPE: Abstract

ABSTRACT: Advanced glycation endproduct (AGE) accumulation in extracellular matrix proteins has been demonstrated in diabetic patients with a significant correlation with the severity of diabetic complications. AGE accumulation induces matrix protein cross-link formation, resulting in an increased stiffness of matrix fibres and the reduction of the susceptibility of matrix proteins to proteolytic degradation. We examined whether glycation-induced collagen cross-linking may affect vascular endothelial cell behaviours such as invasion, proliferation and differentiation, using the in vitro *angiogenesis* model of capillary-like structure formation in three-dimensional matrices of collagen type I. Endothelial cells cultured on collagen gel with angiogenic factors (the combination of fibroblast growth factor-2 and vascular endothelial growth factor) invaded the underlying collagen

LANGUAGE: English

matrix, and organized capillary-like cord structures in the gel. We found that endothelial cell invasion into glycated collagen gel was significantly attenuated without any effect on proteinase activity including cell-associated plasminogen activator and matrix metalloproteinase in the conditioned medium. In addition, subsequent capillary-like cord formation was also *inhibited* in glycated collagen gel. In contrast, endothelial cell proliferation was enhanced on glycated collagen gel with or without angiogenic factors compared with control collagen gel. These results suggest that the structural alterations of extracellular matrix proteins through the glycation-induced cross-link formation affect the interaction between endothelial cell and extracellular matrix, resulting in the impairment of an adequate neovascularization in diabetic patients.

extracellular matrix, resulting in the impairment of an adequate neovascularization in diabetic patients. DESCRIPTORS: MAJOR CONCEPTS: Cardiovascular System (Transport and Circulation) BIOSYSTEMATIC NAMES: Bovidae -- Artiodactyla, Mammalia, Vertebrata, Chordata, Animalia ORGANISMS: bovine (Bovidae) BIOSYSTEMATIC CLASSIFICATION (SUPER TAXA): Animals; Artiodactyls; Chordates; Mammals; Nonhuman Mammals; Nonhuman Vertebrates; Vertebrates DISEASES: diabetes--endocrine disease/pancreas, metabolic disease; vascular endothelial cells advanced glycation endproduct; extracellular CHEMICALS & BIOCHEMICALS: matrix proteins; glycated collagen gel--tissue culture substrate; *type I collagen*--three-dimensional lattices MISCELLANEOUS TERMS: *angiogenesis*; capillary-like structures; collagen-cross linking; neovascularization CONCEPT CODES: Cardiovascular System-General; Methods 14501 Cytology and Cytochemistry-Animal 02506 10060 Biochemical Studies-General 13020 Metabolism-Metabolic Disorders 17002 Endocrine System-General BIOSYSTEMATIC CODES: 85715 Bovidae

7/5/4 (Item 4 from file: 5) DIALOG(R)File 5:Biosis Previews(R) (c) 2002 BIOSIS. All rts. reserv.

11350953 BIOSIS NO.: 199800132285

Three-dimensional type I collagen lattices induce coordinate expression of matrix metalloproteinases MT1-MMP and MMP-2 in microvascular endothelial cells.

AUTHOR: Haas Tara L; Davis Sandra J; Madri Joseph A(a)

AUTHOR ADDRESS: (a) Dep. Pathol., LH115, 310 Cedar St., New Haven, CT 06510 **USA

JOURNAL: Journal of Biological Chemistry 273 (6):p3604-3610 Feb. 6, 1998

ISSN: 0021-9258

DOCUMENT TYPE: Article RECORD TYPE: Abstract LANGUAGE: English

ABSTRACT: Matrix metalloproteinases (MMPs) are hypothesized to play a key role in the processes of endothelial cell migration and matrix remodeling during *angiogenesis*. We utilized an in vitro model of microvascular endothelial cell *angiogenesis*, cells cultured within a collagen matrix, to investigate the MMP profile of endothelial cells undergoing *angiogenesis*. We demonstrated by gelatin zymography that monolayer cultures (two-dimensional) of endothelial cells constitutively expressed low levels of latent MMP-2, but that culture in a three-dimensional collagen matrix increased the total amount of MMP-2 mRNA and protein. Furthermore, 51% of total MMP-2 protein was activated in the three-dimensional culture lysates, compared with 3.5% in two-dimensional culture. The mRNA and protein of MT1-MMP, the putative activator of MMP-2, were up-regulated in endothelial cells cultured in three-dimensional as compared with two-dimensional culture. Treatment of

cultures with MMP *inhibitors* blocked activation of MMP-2 and *inhibited* formation of endothelial cell networks within the collagen gel. Induction of MT1-MMP and MMP-2 appeared to be specific to collagen, inasmuch as culture of the endothelial cells on top of or within, a Matrigel matrix neither increased total MMP-2 nor increased activation of MMP-2. These results suggest that MT1-MMP activation of NMP-2 occurs in endothelial cells undergoing *angiogenesis*, that this activation has a functional role in endothelial cell organization, and that specific matrix interactions may be critical for the increased expression of MT1-MMP and MMP-2.

REGISTRY NUMBERS: 81669-70-7: METALLOPROTEINASE DESCRIPTORS: MAJOR CONCEPTS: Cardiovascular System (Transport and Circulation); Cell Biology; Enzymology (Biochemistry and Molecular Biophysics) BIOSYSTEMATIC NAMES: Muridae--Rodentia, Mammalia, Vertebrata, Chordata, **Animalia** ORGANISMS: rat (Muridae) ORGANISMS: PARTS ETC: microvascular endothelial cell--circulatory system BIOSYSTEMATIC CLASSIFICATION (SUPER TAXA): Animals; Chordates; Mammals; Nonhuman Mammals; Nonhuman Vertebrates; Rodents; Vertebrates CHEMICALS & BIOCHEMICALS: matrix metalloproteinase-2 gene--expression; matrix metalloproteinase-2--activation; membrane-type 1 matrix metalloproteinase gene--expression; membrane-type 1 matrix metalloproteinase--induction; *type I collagen* MISCELLANEOUS TERMS: *angiogenesis*; three-dimensional type I collagen lattice CONCEPT CODES: Cytology and Cytochemistry-Animal 02506 Genetics and Cytogenetics-Animal 03506 10062 Biochemical Studies-Nucleic Acids, Purines and Pyrimidines 10064 Biochemical Studies-Proteins, Peptides and Amino Acids 10808 Enzymes-Physiological Studies 14504 Cardiovascular System-Physiology and Biochemistry BIOSYSTEMATIC CODES: 86375 Muridae

7/5/5 (Item 5 from file: 5)
DIALOG(R)File 5:Biosis Previews(R)
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10530456 BIOSIS NO.: 199699151601

Involvement of the transcription factor NF-kappa-B in tubular morphogenesis of human microvascular endothelial cells by oxidative stress.

AUTHOR: Shono Tadahisa; Ono Mayumi; Izumi Hiroto; Jimi Sei-Ichiro; Matsushima Kouji; Okamoto Takashi; Kohno Kimitoshi; Kuwano Michihiko(a) AUTHOR ADDRESS: (a) Dep. Biochemistry, Kyushu Univ. Sch. Med., Maidashi, Fukuoka 812-82**Japan

JOURNAL: Molecular and Cellular Biology 16 (8):p4231-4239 1996

ISSN: 0270-7306

DOCUMENT TYPE: Article RECORD TYPE: Abstract LANGUAGE: English

ABSTRACT: Oxygen radicals are induced under various pathologic conditions associated with neovascularization. Oxygen radicals modulate *angiogenesis* in cultured human microvascular endothelial cells by an unknown mechanism. Treatment of human microvascular endothelial cells for 15 min with 0.1 to 0.5 mM hydrogen peroxide (H-20-2) or 100 U of tumor necrosis factor alpha per ml induced tubular morphogenesis in type I collagen gels. Gel shift assays with nuclear extracts demonstrated that H-20-2 increases the binding activities of two transcription factors, NF-kappa-B and AP-1, but not of Spl. Tumor necrosis factor alpha increased the binding activities of all three factors. A supershift assay with specific antibodies against JunB, JunD, and c-jun (Jun family) showed that the antibody against c-jun supershifted the AP-1 complex after H-20-2 treatment. Coadministration of the antisense sequence of NF-KB *inhibited* H-20-2-dependent tubular morphogenesis, and the

antisense c-Jun oligonucleotide caused partial *inhibition*. The angiogenic factor responsible for H-20-2-induced tubular morphogenesis was examined. Cellular mRNA levels of vascular endothelial growth factor and interleukin-8 (IL-8), but not those of transforming growth factor et, were increased after treatment with 0.5 mM H-20-2. Coadministration of anti-IL-8 antibody *inhibited* tubular morphogenesis enhanced by H-20-2, and IL-8 itself also enhanced the formation of tube-like structures. Treatment with antisense NF-KB oligonucleotide completely blocked H-20-2-dependent IL-8 production by endothelial cells. The tubular morphogenesis of vascular endothelial cells after treatment with oxidative stimuli and its possible association with NF-kappa-B and IL-8, is examined.

REGISTRY NUMBERS: 7722-84-1: HYDROGEN PEROXIDE DESCRIPTORS: MAJOR CONCEPTS: Biochemistry and Molecular Biophysics; Blood and Lymphatics (Transport and Circulation); Cardiovascular System (Transport and Circulation); Development; Endocrine System (Chemical Coordination and Homeostasis); Genetics; Molecular Genetics (Biochemistry and Molecular Biophysics) BIOSYSTEMATIC NAMES: Hominidae--Primates, Mammalia, Vertebrata, Chordata, Animalia ORGANISMS: Hominidae (Hominidae) BIOSYSTEMATIC CLASSIFICATION (SUPER TAXA): animals; chordates; humans; mammals; primates; vertebrates CHEMICALS & BIOCHEMICALS: HYDROGEN PEROXIDE HYDROGEN PEROXIDE; INTERLEUKIN-8; MESSENGER RNA; MISCELLANEOUS TERMS: NEOVASCULARIZATION; TUMOR NECROSIS FACTOR-ALPHA; *TYPE I COLLAGEN*; VASCULAR ENDOTHELIAL GROWTH FACTOR CONCEPT CODES: 03508 Genetics and Cytogenetics-Human Biochemistry-Gases (1970-) 10012 Replication, Transcription, Translation 10300 Cardiovascular System-Physiology and Biochemistry 14504 Blood, Blood-Forming Organs and Body Fluids-Lymphatic Tissue and 15008 Reticuloendothelial System Endocrine System-General 17002 Developmental Biology-Embryology-Morphogenesis, General 25508 Cytology and Cytochemistry-Human 02508 Biochemical Studies-General 10060 Biochemical Studies-Nucleic Acids, Purines and Pyrimidines 10062 Biochemical Studies-Proteins, Peptides and Amino Acids 10064 Biophysics-Molecular Properties and Macromolecules 10506 BIOSYSTEMATIC CODES: Hominidae 86215 (Item 1 from file: 55)

7/5/6 DIALOG(R) File 55: Biosis Previews (R) (c) 2002 BIOSIS. All rts. reserv.

BIOSIS NO.: 200000012661 12259159

Inhibition by vasoactive intestinal polypeptide (VIP) of *angiogenesis* induced by murine Colon 26-L5 carcinoma cells metastasized in liver.

AUTHOR: Ogasawara Masaru; Murata Jun(a); Kamitani Yukio; Hayashi Kazuko; Saiki Ikuo

AUTHOR ADDRESS: (a) Department of Pathogenic Biochemistry, Institute of Natural Medicine, Toyama Medical and Pharmaceutical University, 2630 Sugitani, Toyama, 930-0194**Japan

JOURNAL: Clinical & Experimental Metastasis 17 (4):p283-291 June, 1999

ISSN: 0262-0898

DOCUMENT TYPE: Article RECORD TYPE: Abstract LANGUAGE: English

SUMMARY LANGUAGE: English

ABSTRACT: We investigated the effect of VIP on the liver metastases and *angiogenesis* by Colon 26-L5 carcinoma cells in mice. Daily systemic administration of VIP, beginning 3 days after tumor inoculation into a portal vein of mice, *inhibited* significantly the development of their liver metastases. Immunohistochemical staining for factor VIII-related antigen in the sections of liver metastases showed that the systemic administration of VIP caused significant prevention of *angiogenesis* within tumor masses. VIP (10-10 to 10-6 M) *inhibited* the invasion of reconstituted basement membrane (Matrigel) by hepatic sinusoidal endothelial (HSE) cells in a concentration-dependent manner in a Transwell chamber assay in vitro and achieved approximately 50% reduction of control at 10-6 M. VIP (10-6 M) also significantly suppressed the haptotactic migration of HSE cells to fibronectin, laminin or type I collagen substrates with a similar *inhibition* rate to the invasion assay. Exposure of VIP to HSE cell s induced accumulation of intracellular cAMP in a concentration-dependent manner. The *inhibitory* effect of VIP (10-6 M) on HSE cell migration was significantly abrogated in the presence of 3 X 10-6 M H-89, a cAMP-dependent protein kinase *inhibitor*. VIP (10-6 M) *inhibited* the morphogenesis of HSE cells into capillary-like structures on Matrigel-coated wells. VIP did not affect the proliferation of HSE cells and the production of gelatinases in HSE cells in vitro at the concentrations used in the invasion assay. These observations suggest that the anti-metastatic effect of VIP on liver metastases by Colon 26-L5 carcinoma cells in mice is partly due to the prevention of tumor *angiogenesis* probably through suppression of the motility of endothelial cells.

REGISTRY NUMBERS: 60-92-4: CYCLIC AMP; 9040-48-6: GELATINASE; 37221-79-7: VASOACTIVE INTESTINAL POLYPEPTIDE DESCRIPTORS: MAJOR CONCEPTS: Digestive System (Ingestion and Assimilation); Tumor Biology BIOSYSTEMATIC NAMES: Muridae--Rodentia, Mammalia, Vertebrata, Chordata, Animalia ORGANISMS: Balb/c mouse (Muridae) -- animal model; colon 26-L5 cell line (Muridae) -- carcinoma cells ORGANISMS: PARTS ETC: hepatic sinusoidal endothelial cells--digestive system, haptotactic migration, morphogenesis, proliferation, tube formation BIOSYSTEMATIC CLASSIFICATION (SUPER TAXA): Animals; Chordates; Mammals; Nonhuman Mammals; Nonhuman Vertebrates; Rodents; Vertebrates DISEASES: colon cancer--digestive system disease, neoplastic disease; liver metastasis--digestive system disease, histopathology, neoplastic disease cAMP {cyclic AMP}--concentration-dependent CHEMICALS & BIOCHEMICALS: intracellular accumulation; fibronectin; gelatinase; laminin; *type I collagen*; vasoactive intestinal polypeptide--anti-angiogenic effect , anti-metastatic effect METHODS & EQUIPMENT: immunohistochemistry--histochemical method tumor *angiogenesis*--*inhibition* MISCELLANEOUS TERMS: ALTERNATE INDEXING: Colonic Neoplasms (MeSH); Liver Neoplasms (MeSH) CONCEPT CODES: Neoplasms and Neoplastic Agents-General 24002 Digestive System-General; Methods BIOSYSTEMATIC CODES: 86375 Muridae

7/5/7 (Item 2 from file: 55)
DIALOG(R)File 55:Biosis Previews(R)
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11974220 BIOSIS NO.: 199900227533

DOCUMENT TYPE: Article

Signaling via fibroblast growth factor receptor-1 is dependent on extracellular matrix in capillary endothelial cell differentiation.

AUTHOR: Kanda Shigeru; Tomasini-Johansson Bianca; Klint Peter; Dixelius Johan; Rubin Kristofer; Claesson-Welsh Lena(a)

AUTHOR ADDRESS: (a) Department of Medical Biochemistry and Microbiology, Biomedical Center, S-751 23, Uppsala**Sweden

JOURNAL: Experimental Cell Research 248 (1):p203-213 April 10, 1999

ISSN: 0014-4827

RECORD TYPE: Abstract LANGUAGE: English

SUMMARY LANGUAGE: English

ABSTRACT: Differentiation of endothelial cells, i.e., formation of a vessel lumen, is a prerequisite for *angiogenesis*. The underlying molecular mechanisms are ill defined. We have studied a brain capillary endothelial cell line (IBEC) established from H-2Kb-tsA58 transgenic mice. These cells form hollow tubes in three-dimensional type I collagen gels in response to fibroblast growth factor-2 (FGF-2). Culture of IBEC on collagen gels in the presence of FGF-2 protected cells from apoptosis and allowed tube formation (i.e., differentiation) but not growth of the cells. FGF-induced differentiation, but not cell survival, was *inhibited* by treatment of the cells with an anti-betal-integrin IgG. Changes in integrin expression in the collagen-gel cultures could not be detected. Rather, cell-matrix interactions critical for endothelial cell differentiation were created during the culture, as indicated by the gradual increase in tyrosine phosphorylation of focal adhesion kinase in the collagen-gel cultures. Inclusion of laminin in the collagen gels led to FGF-2-independent formation of tube structures, but cells were not protected from apoptosis. These data indicate that FGF receptor-1 signal transduction in this cell model results in cell survival. Through mechanisms dependent on cell-matrix interactions, possibly involving the alpha3beta1-integrin and laminin produced by the collagen-cultured IBE cells, FGF stimulation also leads to differentiation of the cells.

REGISTRY NUMBERS: 60-18-4Q: TYROSINE; 556-03-6Q: TYROSINE; 9031-44-1: KINASE **DESCRIPTORS:** MAJOR CONCEPTS: Cardiovascular System (Transport and Circulation); Cell BIOSYSTEMATIC NAMES: Muridae--Rodentia, Mammalia, Vertebrata, Chordata, ORGANISMS: IBEC cell line (Muridae) -- brain capillary endothelial cell ORGANISMS: PARTS ETC: capillary endothelial cell--circulatory system, differentiation BIOSYSTEMATIC CLASSIFICATION (SUPER TAXA): Animals; Chordates; Mammals; Nonhuman Mammals; Nonhuman Vertebrates; Rodents; Vertebrates extracellular matrix; fibroblast growth CHEMICALS & BIOCHEMICALS: factor receptor-1; fibroblast growth factor-2; focal adhesion kinase --tyrosine phosphorylation; *type I collagen*; tyrosine MISCELLANEOUS TERMS: *angiogenesis*; signal transduction CONCEPT CODES: 02506 Cytology and Cytochemistry-Animal 10060 Biochemical Studies-General Enzymes-General and Comparative Studies; Coenzymes 10802 Cardiovascular System-General; Methods 14501 Endocrine System-General 17002 20501 Nervous System-General; Methods BIOSYSTEMATIC CODES: 86375 Muridae 7/5/8 (Item 3 from file: 55) DIALOG(R) File 55: Biosis Previews(R) (c) 2002 BIOSIS. All rts. reserv.

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11486517 BIOSIS NO.: 199800267849

Inhibition of *angiogenesis* on glycated collagen lattices.

AUTHOR: Kuzuya M(a); Satake S; Ai S; Asai T; Kanda S; Ramos M A; Miura H; Ueda M; Iguchi A

AUTHOR ADDRESS: (a)Dep. Geriatr., Nagoya Univ. Sch. Med., 65 Tsuruma-cho, Showa-ku, Nagoya 466-8550**Japan

JOURNAL: Diabetologia 41 (5):p491-499 May, 1998

ISSN: 0012-186X

DOCUMENT TYPE: Article

RECORD TYPE: Abstract

LANGUAGE: English

ABSTRACT: Advanced glycation endproduct (AGE) accumulation in extracellular matrix proteins has been demonstrated in diabetic patients with a significant correlation with the severity of diabetic complications. AGE accumulation induces matrix protein cross-link formation, resulting in an increased stiffness of matrix fibres and the reduction of the susceptibility of matrix proteins to proteolytic degradation. We examined whether glycation-induced collagen cross-linking may affect vascular endothelial cell behaviours such as invasion, proliferation and differentiation, using the in vitro *angiogenesis* model of capillary-like structure formation in three-dimensional matrices of collagen type I. Endothelial cells cultured on collagen gel with angiogenic factors (the combination of fibroblast growth factor-2 and vascular endothelial growth factor) invaded the underlying collagen matrix, and organized capillary-like cord structures in the gel. We found that endothelial cell invasion into glycated collagen gel was significantly attenuated without any effect on proteinase activity including cell-associated plasminogen activator and matrix metalloproteinase in the conditioned medium. In addition, subsequent capillary-like cord formation was also *inhibited* in glycated collagen gel. In contrast, endothelial cell proliferation was enhanced on glycated collagen gel with or without angiogenic factors compared with control collagen gel. These results suggest that the structural alterations of extracellular matrix proteins through the glycation-induced cross-link formation affect the interaction between endothelial cell and extracellular matrix, resulting in the impairment of an adequate neovascularization in diabetic patients. DESCRIPTORS: MAJOR CONCEPTS: Cardiovascular System (Transport and Circulation) BIOSYSTEMATIC NAMES: Bovidae--Artiodactyla, Mammalia, Vertebrata, Chordata, Animalia ORGANISMS: bovine (Bovidae) BIOSYSTEMATIC CLASSIFICATION (SUPER TAXA): Animals; Artiodactyls; Chordates; Mammals; Nonhuman Mammals; Nonhuman Vertebrates; Vertebrates DISEASES: diabetes -- endocrine disease/pancreas, metabolic disease; vascular endothelial cells advanced glycation endproduct; extracellular CHEMICALS & BIOCHEMICALS: matrix proteins; glycated collagen gel--tissue culture substrate; *type I collagen*--three-dimensional lattices *angiogenesis*; capillary-like structures; MISCELLANEOUS TERMS: collagen-cross linking; neovascularization CONCEPT CODES: 14501 Cardiovascular System-General; Methods Cytology and Cytochemistry-Animal 02506 Biochemical Studies-General 10060 13020 Metabolism-Metabolic Disorders 17002 Endocrine System-General BIOSYSTEMATIC CODES: 85715 Bovidae (Item 4 from file: 55) DIALOG(R) File 55: Biosis Previews(R)

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BIOSIS NO.: 199800132285

Three-dimensional type I collagen lattices induce coordinate expression of matrix metalloproteinases MT1-MMP and MMP-2 in microvascular endothelial cells.

AUTHOR: Haas Tara L; Davis Sandra J; Madri Joseph A(a)

AUTHOR ADDRESS: (a) Dep. Pathol., LH115, 310 Cedar St., New Haven, CT 06510

JOURNAL: Journal of Biological Chemistry 273 (6):p3604-3610 Feb. 6, 1998

ISSN: 0021-9258

DOCUMENT TYPE: Article RECORD TYPE: Abstract LANGUAGE: English

ABSTRACT: Matrix metalloproteinases (MMPs) are hypothesized to play a key

role in the processes of endothelial cell migration and matrix remodeling during *angiogenesis*. We utilized an in vitro model of microvascular endothelial cell *angiogenesis*, cells cultured within a collagen matrix, to investigate the MMP profile of endothelial cells undergoing *angiogenesis*. We demonstrated by gelatin zymography that monolayer cultures (two-dimensional) of endothelial cells constitutively expressed low levels of latent MMP-2, but that culture in a three-dimensional collagen matrix increased the total amount of MMP-2 mRNA and protein. Furthermore, 51% of total MMP-2 protein was activated in the three-dimensional culture lysates, compared with 3.5% in two-dimensional culture. The mRNA and protein of MT1-MMP, the putative activator of MMP-2, were up-regulated in endothelial cells cultured in three-dimensional as compared with two-dimensional culture. Treatment of cultures with MMP *inhibitors* blocked activation of MMP-2 and *inhibited* formation of endothelial cell networks within the collagen gel. Induction of MT1-MMP and MMP-2 appeared to be specific to collagen, inasmuch as culture of the endothelial cells on top of or within, a Matrigel matrix neither increased total MMP-2 nor increased activation of MMP-2. These results suggest that MT1-MMP activation of NMP-2 occurs in endothelial cells undergoing *angiogenesis*, that this activation has a functional role in endothelial cell organization, and that specific matrix interactions may be critical for the increased expression of MT1-MMP and MMP-2.

REGISTRY NUMBERS: 81669-70-7: METALLOPROTEINASE DESCRIPTORS: MAJOR CONCEPTS: Cardiovascular System (Transport and Circulation); Cell Biology; Enzymology (Biochemistry and Molecular Biophysics) BIOSYSTEMATIC NAMES: Muridae--Rodentia, Mammalia, Vertebrata, Chordata, Animalia ORGANISMS: rat (Muridae) ORGANISMS: PARTS ETC: microvascular endothelial cell--circulatory system BIOSYSTEMATIC CLASSIFICATION (SUPER TAXA): Animals; Chordates; Mammals; Nonhuman Mammals; Nonhuman Vertebrates; Rodents; Vertebrates CHEMICALS & BIOCHEMICALS: matrix metalloproteinase-2 gene--expression; matrix metalloproteinase-2--activation; membrane-type 1 matrix metalloproteinase gene--expression; membrane-type 1 matrix metalloproteinase--induction; *type I collagen* MISCELLANEOUS TERMS: *angiogenesis*; three-dimensional type I collagen lattice CONCEPT CODES: Cytology and Cytochemistry-Animal 02506 Genetics and Cytogenetics-Animal 03506 Biochemical Studies-Nucleic Acids, Purines and Pyrimidines 10062 10064 Biochemical Studies-Proteins, Peptides and Amino Acids 10808 Enzymes-Physiological Studies 14504 Cardiovascular System-Physiology and Biochemistry BIOSYSTEMATIC CODES: 86375 Muridae

7/5/10 (Item 5 from file: 55)
DIALOG(R)File 55:Biosis Previews(R)
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10530456 BIOSIS NO.: 199699151601

Involvement of the transcription factor NF-kappa-B in tubular morphogenesis of human microvascular endothelial cells by oxidative stress.

AUTHOR: Shono Tadahisa; Ono Mayumi; Izumi Hiroto; Jimi Sei-Ichiro; Matsushima Kouji; Okamoto Takashi; Kohno Kimitoshi; Kuwano Michihiko(a) AUTHOR ADDRESS: (a) Dep. Biochemistry, Kyushu Univ. Sch. Med., Maidashi, Fukuoka 812-82**Japan

JOURNAL: Molecular and Cellular Biology 16 (8):p4231-4239 1996

ISSN: 0270-7306

DOCUMENT TYPE: Article RECORD TYPE: Abstract LANGUAGE: English

ABSTRACT: Oxygen radicals are induced under various pathologic conditions

associated with neovascularization. Oxygen radicals modulate *angiogenesis* in cultured human microvascular endothelial cells by an unknown mechanism. Treatment of human microvascular endothelial cells for 15 min with 0.1 to 0.5 mM hydrogen peroxide (H-20-2) or 100 U of tumor necrosis factor alpha per ml induced tubular morphogenesis in type I collagen gels. Gel shift assays with nuclear extracts demonstrated that H-20-2 increases the binding activities of two transcription factors, NF-kappa-B and AP-1, but not of Spl. Tumor necrosis factor alpha increased the binding activities of all three factors. A supershift assay with specific antibodies against JunB, JunD, and c-jun (Jun family) showed that the antibody against c-jun supershifted the AP-1 complex after H-20-2 treatment. Coadministration of the antisense sequence of NF-KB *inhibited* H-20-2-dependent tubular morphogenesis, and the antisense c-Jun oligonucleotide caused partial *inhibition*. The angiogenic factor responsible for H-20-2-induced tubular morphogenesis was examined. Cellular mRNA levels of vascular endothelial growth factor and interleukin-8 (IL-8), but not those of transforming growth factor et, were increased after treatment with 0.5 mM H-20-2. Coadministration of anti-IL-8 antibody *inhibited* tubular morphogenesis enhanced by H-20-2, and IL-8 itself also enhanced the formation of tube-like structures. Treatment with antisense NF-KB oligonucleotide completely blocked H-20-2-dependent IL-8 production by endothelial cells. The tubular morphogenesis of vascular endothelial cells after treatment with oxidative stimuli and its possible association with NF-kappa-B and IL-8,

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is examined.
REGISTRY NUMBERS: 7722-84-1: HYDROGEN PEROXIDE
DESCRIPTORS:
 MAJOR CONCEPTS: Biochemistry and Molecular Biophysics; Blood and
    Lymphatics (Transport and Circulation); Cardiovascular System
    (Transport and Circulation); Development; Endocrine System (Chemical
    Coordination and Homeostasis); Genetics; Molecular Genetics
    (Biochemistry and Molecular Biophysics)
  BIOSYSTEMATIC NAMES: Hominidae--Primates, Mammalia, Vertebrata, Chordata,
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  ORGANISMS: Hominidae (Hominidae)
  BIOSYSTEMATIC CLASSIFICATION (SUPER TAXA): animals; chordates; humans;
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                              HYDROGEN PEROXIDE
  CHEMICALS & BIOCHEMICALS:
  MISCELLANEOUS TERMS: HYDROGEN PEROXIDE; INTERLEUKIN-8; MESSENGER RNA;
    NEOVASCULARIZATION; TUMOR NECROSIS FACTOR-ALPHA; *TYPE I COLLAGEN*;
    VASCULAR ENDOTHELIAL GROWTH FACTOR
CONCEPT CODES:
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          Biochemistry-Gases (1970-)
  10012
          Replication, Transcription, Translation
  10300
          Cardiovascular System-Physiology and Biochemistry
  14504
          Blood, Blood-Forming Organs and Body Fluids-Lymphatic Tissue and
  15008
             Reticuloendothelial System
          Endocrine System-General
  17002
          Developmental Biology-Embryology-Morphogenesis, General
  25508
          Cytology and Cytochemistry-Human
  02508
          Biochemical Studies-General
  10060
          Biochemical Studies-Nucleic Acids, Purines and Pyrimidines
  10062
          Biochemical Studies-Proteins, Peptides and Amino Acids
  10064
          Biophysics-Molecular Properties and Macromolecules
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BIOSYSTEMATIC CODES:
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DIALOG(R) File 143:Biol. & Agric. Index
(c) 2002 The HW Wilson Co. All rts. reserv.
           H.W. WILSON RECORD NUMBER: BBA198012778
Disruption of angiogenesis by PEX, a noncatalytic metalloproteinase
  fragment with integrin binding activity
 *Brooks, Peter C*
 Silletti, Steve; von Schalscha, Tami L
 Cell v. 92 (Feb. 6 1998) p. 391-400
                                   ISSN: 0092-8674 LANGUAGE:
 DOCUMENT TYPE: Feature Article
 RECORD STATUS: Corrected or revised record
 DESCRIPTORS: Integrins; Angiogenesis inhibitors; Metalloproteinases
            (Item 2 from file: 143)
DIALOG(R) File 143:Biol. & Agric. Index
(c) 2002 The HW Wilson Co. All rts. reserv.
           H.W. WILSON RECORD NUMBER: BBAI00060824
Contact with fibrillar collagen inhibits melanoma cell proliferation by
  up-regulating p27KIP1
 Henriet, Patrick
 Zhong, Zhi-Duan; *Brooks, Peter C*
 Proceedings of the National Academy of Sciences of the United States of
America v. 97 no18 (Aug. 29 2000) p. 10026-31
                                   ISSN: 0027-8424 LANGUAGE: English
 DOCUMENT TYPE: Feature Article
 RECORD STATUS: Corrected or revised record
 DESCRIPTORS: Collagen; Melanoma; Cell proliferation -- Inhibition
            (Item 3 from file: 143)
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DIALOG(R) File 143: Biol. & Agric. Index
(c) 2002 The HW Wilson Co. All rts. reserv.
           H.W. WILSON RECORD NUMBER: BBA195002655
Integrin avb3 antagonists promote tumor regression by inducing apoptosis of
  angiogenic blood vessels
 *Brooks, Peter C*
 Montgomery, Anthony M. P; Rosenfeld, Mauricio
 Cell v. 79 (Dec. 30 1994) p. 1157-64
                                    ISSN: 0092-8674 LANGUAGE: English
 DOCUMENT TYPE: Feature Article
 RECORD STATUS: Corrected or revised record
 DESCRIPTORS: Angiogenesis; Integrins; Tumor cells--Blood supply
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DIALOG(R) File 143: Biol. & Agric. Index
(c) 2002 The HW Wilson Co. All rts. reserv.
          H.W. WILSON RECORD NUMBER: BBAI96028591
Localization of matrix metalloproteinase MMP-2 to the surface of invasive
  cells by interaction with integrin avb3
 *Brooks, Peter C*
 Stromblad, Staffan; Sanders, Luraynne C
 Cell v. 85 (May 31 '96) p. 683-93
                                   ISSN: 0092-8674 LANGUAGE:
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 DOCUMENT TYPE: Feature Article
 RECORD STATUS: Corrected or revised record
 DESCRIPTORS: Gelatinase A; Integrins
            (Item 5 from file: 143)
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DIALOG(R) File 143: Biol. & Agric. Index
(c) 2002 The HW Wilson Co. All rts. reserv.
          H.W. WILSON RECORD NUMBER: BBA195030391
Requirement of the NPXY motif in the integrin b3 subunit cytoplasmic tail
  for melanoma cell migration in vitro and in vivo
 Filardo, Edward J
 *Brooks, Peter C*; Deming, Sandra L
 The Journal of Cell Biology v. 130 no2 (July '95) p. 441-50
                                   ISSN: 0021-9525 LANGUAGE: English
 DOCUMENT TYPE: Feature Article
 RECORD STATUS: Corrected or revised record
 DESCRIPTORS: Integrins; Melanoma; Tumor cell migration
            (Item 6 from file: 143)
DIALOG(R) File 143: Biol. & Agric. Index
(c) 2002 The HW Wilson Co. All rts. reserv.
          H.W. WILSON RECORD NUMBER: BBA193045451
Subtractive immunization yields monoclonal antibodies that specifically
  inhibit metastasis
 *Brooks, Peter C*
 Lin, Jian-Min; French, Deborah L
 The Journal of Cell Biology v. 122 no6 (Sept. '93) p. 1351-9
 DOCUMENT TYPE: Feature Article ISSN: 0021-9525 LANGUAGE:
                                                               English
 RECORD STATUS: New record
 DESCRIPTORS: Metastasis; Vaccines and vaccination; Monoclonal antibodies
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DIALOG(R) File 143: Biol. & Agric. Index
(c) 2002 The HW Wilson Co. All rts. reserv.
           H.W. WILSON RECORD NUMBER: BBAI01052325
Proteolytic exposure of a cryptic site within collage type IV is required
  for angiogenesis and tumor growth in vivo
 Xu, Jingsong
 Rodriguez, Dorothy; *Petitclerc, Eric*
 The Journal of Cell Biology v. 154 no5 (Sept. 3 2001) p. 1069-79
                                  ISSN: 0021-9525 LANGUAGE: English
 DOCUMENT TYPE: Feature Article
 RECORD STATUS: Corrected or revised record
 In: The Journal of Cell Biology v. 154 no5 (Sept. 3 2001) p. 1069-79;
Correction. v155 no5 p859 N 26 2001.
 DESCRIPTORS: Gelatinase A; Angiogenesis; Proteolysis
            (Item 2 from file: 143)
 13/5/2
DIALOG(R) File 143: Biol. & Agric. Index
(c) 2002 The HW Wilson Co. All rts. reserv.
           H.W. WILSON RECORD NUMBER: BBAI01050691
Plasminogen activator inhibitor-1 regulates tumor growth and angiogenesis
 McMahon, Grainne A
 *Petitclerc, Eric*; Stefansson, Steingrimur
 The Journal of Biological Chemistry v. 276 no36 (Sept. 7 2001) p. 33964-8
 DOCUMENT TYPE: Feature Article ISSN: 0021-9258 LANGUAGE: English
 RECORD STATUS: New record
 DESCRIPTORS: Plasminogen activator inhibitors; Angiogenesis inhibitors
            (Item 3 from file: 143)
 13/5/3
DIALOG(R) File 143: Biol. & Agric. Index
(c) 2002 The HW Wilson Co. All rts. reserv.
           H.W. WILSON RECORD NUMBER: BBAI01016615
1357200
Inhibition of angiogenesis in vivo by plasminogen activator inhibitor-1
 Stefansson, Steingrimur
 *Petitclerc, Eric*; Wong, Michael K. K
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The Journal of Biological Chemistry v. 276 no11 (Mar. 16 2001) p. 8135-41
 DOCUMENT TYPE: Feature Article ISSN: 0021-9258 LANGUAGE: English
 RECORD STATUS: New record
 DESCRIPTORS: Angiogenesis inhibitors; Plasminogen activator inhibitors
            (Item 4 from file: 143)
 13/5/4
DIALOG(R) File 143: Biol. & Agric. Index
(c) 2002 The HW Wilson Co. All rts. reserv.
          H.W. WILSON RECORD NUMBER: BBAI00024786
1297247
New functions for non-collagenous domains of human collagen type IV. Novel
  integrin ligands inhibiting angiogenesis and tumor growth in vivo
 *Petitclerc, Eric*
 Boutaud, Ariel; Prestayko, Archie
 The Journal of Biological Chemistry v. 275 no11 (Mar. 17 2000) p. 8051-61
 DOCUMENT TYPE: Feature Article ISSN: 0021-9258 LANGUAGE: English
 RECORD STATUS: Corrected or revised record
 DESCRIPTORS: Collagen; Integrins; Tumor cells--Blood supply; Angiogenesis
  inhibitors
 13/5/5
            (Item 5 from file: 143)
DIALOG(R) File 143: Biol. & Agric. Index
(c) 2002 The HW Wilson Co. All rts. reserv.
          H.W. WILSON RECORD NUMBER: BBA197015381
Mechanisms of action of antimalarials in inflammation. Induction of
  apoptosis in human endothelial cells
 Potvin, Frederic
 *Petitclerc, Eric*; Marceau, Francois
 Journal of Immunology v. 158 (Feb. 15 '97) p. 1872-9
 DOCUMENT TYPE: Feature Article ISSN: 0022-1767 LANGUAGE: English
 RECORD STATUS: New record
 DESCRIPTORS: Apoptosis--Man; Antimalarials; Inflammation--Man
            (Item 6 from file: 143)
DIALOG(R) File 143:Biol. & Agric. Index
(c) 2002 The HW Wilson Co. All rts. reserv.
           H.W. WILSON RECORD NUMBER: BBAI96021621
Pathologic leukocyte infiltration of the rabbit aorta confers a vasomotor
  effect to chemotactic peptides through cyclooxygenase-derived metabolites
 *Petitclerc, Eric*
 Levesque, Luc; Grose, John H
 Journal of Immunology v. 156 (May 1 '96) p. 3426-34
 DOCUMENT TYPE: Feature Article ISSN: 0022-1767 LANGUAGE: English
 RECORD STATUS: New record
 DESCRIPTORS: Leukocytes; Vasomotor system--Physiology; Metabolites;
  Chemotactic factors
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(Item 1 from file: 143) 15/5/1 DIALOG(R) File 143: Biol. & Agric. Index (c) 2002 The HW Wilson Co. All rts. reserv.

H.W. WILSON RECORD NUMBER: BBA196029523 1476719

Graded activation of fibroblast growth factor receptor 3 by mutations causing achondroplasia and thanatophoric dysplasia

Naski, Michael C

Wang, Qing; *Xu, Jingsong*

Nature Genetics v. 13 (June 1996) p. 233-7

ISSN: 1061-4036 LANGUAGE: English DOCUMENT TYPE: Feature Article

RECORD STATUS: Corrected or revised record

DESCRIPTORS: Fibroblast growth factor receptors--Man; Achondroplasia--Man ; Mutation (Biology) -- Man

(Item 2 from file: 143) 15/5/2 DIALOG(R) File 143: Biol. & Agric. Index (c) 2002 The HW Wilson Co. All rts. reserv.

H.W. WILSON RECORD NUMBER: BBAI01052325

Proteolytic exposure of a cryptic site within collage type IV is required for angiogenesis and tumor growth in vivo

Xu, Jingsong

Rodriguez, Dorothy; Petitclerc, Eric

The Journal of Cell Biology v. 154 no5 (Sept. 3 2001) p. 1069-79

ISSN: 0021-9525 LANGUAGE: English DOCUMENT TYPE: Feature Article

RECORD STATUS: Corrected or revised record

In: The Journal of Cell Biology v. 154 no5 (Sept. 3 2001) p. 1069-79; Correction. v155 no5 p859 N 26 2001.

DESCRIPTORS: Gelatinase A; Angiogenesis; Proteolysis

(Item 3 from file: 143) DIALOG(R) File 143: Biol. & Agric. Index (c) 2002 The HW Wilson Co. All rts. reserv.

H.W. WILSON RECORD NUMBER: BBAI00034250

Temporal and spatial gradients of Fgf8 and Fgf17 regulate proliferation and differentiation of midline cerebellar structures

Xu, Jingsong

Liu, Zhonghao; Ornitz, David M

Development (Cambridge, England) v. 127 no9 (May 2000) p. 1833-43

DOCUMENT TYPE: Feature Article ISSN: 0950-1991 LANGUAGE: English

RECORD STATUS: New record

DESCRIPTORS: Organogenesis -- Cerebellum; Developmental genetics; Neurogenesis; Fibroblast growth factor

(Item 4 from file: 143) 15/5/4 DIALOG(R) File 143:Biol. & Agric. Index (c) 2002 The HW Wilson Co. All rts. reserv. 0879363 H.W. WILSON RECORD NUMBER: BBAI98046483 Identification of the cytoplasmic regions of fibroblast growth factor (FGF) receptor 1 which play important roles in induction of neurite outgrowth in PC12 cells by FGF-1 Lin, Hsien-Yi *Xu, Jingsong*; Ischenko, Irene Molecular and Cellular Biology v. 18 no7 (July '98) p. 3762-70 ISSN: 0270-7306 LANGUAGE: English DOCUMENT TYPE: Feature Article RECORD STATUS: New record DESCRIPTORS: Fibroblast growth factor receptors; Neurogenesis (Item 5 from file: 143) 15/5/5 DIALOG(R) File 143: Biol. & Agric. Index (c) 2002 The HW Wilson Co. All rts. reserv. H.W. WILSON RECORD NUMBER: BBAI98004907 0801407 Transplanted oligodendrocyte progenitor cells expressing a dominant-negative FGF receptor transgene fail to migrate in vivo Osterhout, Donna J Ebner, Sylvie; *Xu, Jingsong* Journal of Neuroscience v. 17 (Dec. 1 '97) p. 9122-32 DOCUMENT TYPE: Feature Article ISSN: 0270-6474 LANGUAGE: English RECORD STATUS: New record DESCRIPTORS: Fibroblast growth factor receptors; Neuroglia; Neurogenesis; Cell migration 15/5/6 (Item 6 from file: 143) DIALOG(R) File 143:Biol. & Agric. Index (c) 2002 The HW Wilson Co. All rts. reserv. H.W. WILSON RECORD NUMBER: BBA196039623 The fibroblast growth factor receptor-1 is necessary for the induction of neurite outgrowth in PC12 cells by aFGF Lin, Hsien-Yi *Xu, Jingsong*; Ornitz, David M Journal of Neuroscience v. 16 (Aug. 1 '96) p. 4579-87 DOCUMENT TYPE: Feature Article ISSN: 0270-6474 LANGUAGE: English RECORD STATUS: New record DESCRIPTORS: Differentiation (Biology); Fibroblast growth factor receptors (Item 7 from file: 143) DIALOG(R) File 143:Biol. & Agric. Index (c) 2002 The HW Wilson Co. All rts. reserv. 0601977 H.W. WILSON RECORD NUMBER: BBAI96034385 Receptor specificity of the fibroblast growth factor family Ornitz, David M *Xu, Jingsong*; Colvin, Jennifer S The Journal of Biological Chemistry v. 271 (June 21 '96) p. 15292-7 DOCUMENT TYPE: Feature Article ISSN: 0021-9258 LANGUAGE: English RECORD STATUS: New record

15/5/8 (Item 8 from file: 143)

DESCRIPTORS: Fibroblast growth factor receptors

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DIALOG(R) File 143:Biol. & Agric. Index
(c) 2002 The HW Wilson Co. All rts. reserv.
           H.W. WILSON RECORD NUMBER: BBAI96001844
0559124
FGF-8 isoforms activate receptor splice forms that are expressed in
 mesenchymal regions of mouse development
MacArthur, Craig A
 Lawshe, Avril; *Xu, Jingsong*
Development (Cambridge, England) v. 121 (Nov. '95) p. 3603-13
DOCUMENT TYPE: Feature Article
                                  ISSN: 0950-1991 LANGUAGE: English
 RECORD STATUS: New record
DESCRIPTORS: Pattern (Biology); Mesenchyme; Developmental genetics;
  Fibroblast growth factor
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10530456 BIOSIS NO.: 199699151601

Involvement of the transcription factor NF-kappa-B in tubular morphogenesis of human microvascular endothelial cells by oxidative stress.

AUTHOR: Shono Tadahisa; Ono Mayumi; Izumi Hiroto; Jimi Sei-Ichiro; Matsushima Kouji; Okamoto Takashi; Kohno Kimitoshi; Kuwano Michihiko(a) AUTHOR ADDRESS: (a) Dep. Biochemistry, Kyushu Univ. Sch. Med., Maidashi, Fukuoka 812-82**Japan

JOURNAL: Molecular and Cellular Biology 16 (8):p4231-4239 1996

ISSN: 0270-7306

DOCUMENT TYPE: Article RECORD TYPE: Abstract LANGUAGE: English

ABSTRACT: Oxygen radicals are induced under various pathologic conditions associated with neovascularization. Oxygen radicals modulate *angiogenesis* in cultured human microvascular endothelial cells by an unknown mechanism. Treatment of human microvascular endothelial cells for 15 min with 0.1 to 0.5 mM hydrogen peroxide (H-20-2) or 100 U of tumor necrosis factor alpha per ml induced tubular morphogenesis in type I collagen gels. Gel shift assays with nuclear extracts demonstrated that H-20-2 increases the binding activities of two transcription factors, NF-kappa-B and AP-1, but not of Spl. Tumor necrosis factor alpha increased the binding activities of all three factors. A supershift assay with specific *antibodies* against JunB, JunD, and c-jun (Jun family) showed that the *antibody* against c-jun supershifted the AP-1 complex after H-20-2 treatment. Coadministration of the antisense sequence of NF-KB *inhibited* H-20-2-dependent tubular morphogenesis, and the antisense c-Jun oligonucleotide caused partial *inhibition*. The angiogenic factor responsible for H-20-2-induced tubular morphogenesis was examined. Cellular mRNA levels of vascular endothelial growth factor and interleukin-8 (IL-8), but not those of transforming growth factor et, were increased after treatment with 0.5 mM H-20-2. Coadministration of anti-IL-8 *antibody* *inhibited* tubular morphogenesis enhanced by H-20-2, and IL-8 itself also enhanced the formation of tube-like structures. Treatment with antisense NF-KB oligonucleotide completely blocked H-20-2-dependent IL-8 production by endothelial cells. The tubular morphogenesis of vascular endothelial cells after treatment with oxidative stimuli and its possible association with NF-kappa-B and IL-8, is examined.

REGISTRY NUMBERS: 7722-84-1: HYDROGEN PEROXIDE DESCRIPTORS:

MAJOR CONCEPTS: Biochemistry and Molecular Biophysics; Blood and Lymphatics (Transport and Circulation); Cardiovascular System (Transport and Circulation); Development; Endocrine System (Chemical Coordination and Homeostasis); Genetics; Molecular Genetics (Biochemistry and Molecular Biophysics)

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BIOSYSTEMATIC NAMES: Hominidae--Primates, Mammalia, Vertebrata, Chordata,
   Animalia
  ORGANISMS: Hominidae (Hominidae)
  BIOSYSTEMATIC CLASSIFICATION (SUPER TAXA): animals; chordates; humans;
    mammals; primates; vertebrates
  CHEMICALS & BIOCHEMICALS:
                              HYDROGEN PEROXIDE
                         HYDROGEN PEROXIDE; INTERLEUKIN-8; MESSENGER RNA;
  MISCELLANEOUS TERMS:
   NEOVASCULARIZATION; TUMOR NECROSIS FACTOR-ALPHA; *TYPE I COLLAGEN*;
    VASCULAR ENDOTHELIAL GROWTH FACTOR
CONCEPT CODES:
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          Genetics and Cytogenetics-Human
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          Biochemistry-Gases (1970-)
          Replication, Transcription, Translation
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          Cardiovascular System-Physiology and Biochemistry
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          Blood, Blood-Forming Organs and Body Fluids-Lymphatic Tissue and
  15008
             Reticuloendothelial System
          Endocrine System-General
  17002
          Developmental Biology-Embryology-Morphogenesis, General
  25508
  02508
          Cytology and Cytochemistry-Human
          Biochemical Studies-General
  10060
          Biochemical Studies-Nucleic Acids, Purines and Pyrimidines
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ISSN: 0945-053X

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DOCUMENT TYPE: Article RECORD TYPE: Abstract LANGUAGE: English

ABSTRACT: Prostate *cancer* is the second leading cause of male *cancer* -related deaths in the United States. Interestingly, prostate *cancer* preferentially metastasizes to bone. Once in the bone microenvironment, advanced prostate *cancer* becomes highly resistant to therapeutic modalities. Several factors, such as, extracelluar matrix components, have been implicated in the spread and propagation of prostatic carcinoma. The prostate cell line, PC3, adhere and spread on collagen I to a greater degree than on fibronectin (FN) or poly-L-lysine (PLL). Flow cytometry analysis reveals the presence of the alpha1, alpha2 and alpha3 collagen binding integrin subunits. *Antibody* function blocking studies reveal that PC3 cells can utilize alpha2beta1 and alpha3beta1 integrins to adhere to collagen I. Cells plated on collagen I exhibit increased rates of proliferation over cells plated on FN or tissue culture plastic. Additionally, cells plated on collagen I show increased expression of cyclin D1, a molecule associated with progression through G1 phase of the cell cycle. *Inhibitor* studies point to a role for phosphatidylinositol 3-kinase (PI3K), map kinase (MAPK) and p70 S6 kinase in collagen I-mediated PC3 cell proliferation and cyclin D1 expression. Type I collagen may facilitate the colonization and growth of metastatic prostate tumor cells in the bone microenvironment.

REGISTRY NUMBERS: 142243-02-5: MITOGEN-ACTIVATED PROTEIN KINASE; 115926-52-8: PHOSPHATIDYLINOSITOL 3-KINASE; 25104-18-1Q: POLY-L-LYSINE; 38000-06-5Q: POLY-L-LYSINE DESCRIPTORS: MAJOR CONCEPTS: Enzymology (Biochemistry and Molecular Biophysics); Reproductive System (Reproduction); Skeletal System (Movement and Support); Tumor Biology BIOSYSTEMATIC NAMES: Hominidae--Primates, Mammalia, Vertebrata, Chordata, Animalia ORGANISMS: PC3 cell line (Hominidae) -- human prostate carcinoma cells ORGANISMS: PARTS ETC: bone--skeletal system BIOSYSTEMATIC CLASSIFICATION (SUPER TAXA): Animals; Chordates; Humans; Mammals; Primates; Vertebrates DISEASES: bone metastasis--bone disease, neoplastic disease; prostate *cancer*--neoplastic disease, reproductive system disease/male, urologic disease alpha-1-collagen binding integrin subunit; CHEMICALS & BIOCHEMICALS: alpha-2-collagen binding integrin subunit; alpha-3-collagen binding integrin subunit; cyclin D1; extracellular matrix components; fibronectin; mitogen-activated protein kinase {MAPK}; p70 S6 kinase; phosphatidylinositol 3-kinase {PI3K}; poly-L-lysine; *type I collagen* METHODS & EQUIPMENT: *antibody* function blocking study--analytical method; flow cytometry--cytological method, cytophotometry MISCELLANEOUS TERMS: cell proliferation ALTERNATE INDEXING: Bone Neoplasms (MeSH); Prostatic Neoplasms (MeSH) CONCEPT CODES: 02508 Cytology and Cytochemistry-Human Biochemical Studies-Proteins, Peptides and Amino Acids 10064 Enzymes-General and Comparative Studies; Coenzymes 10802 Urinary System and External Secretions-Pathology 15506

Reproductive System-Physiology and Biochemistry

Bones, Joints, Fasciae, Connective and Adipose Tissue-Physiology

Bones, Joints, Fasciae, Connective and Adipose Tissue-Pathology

Neoplasms and Neoplastic Agents-Pathology; Clinical Aspects;

Reproductive System-Pathology

and Biochemistry

Systemic Effects
BIOSYSTEMATIC CODES:
86215 Hominidae

22/5/2 (Item 2 from file: 5)
DIALOG(R)File 5:Biosis Previews(R)
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13207759 BIOSIS NO.: 200100414908

Functional interplay between type I collagen and cell surface matrix metalloproteinase activity.

AUTHOR: Ellerbroek Shawn M; Wu Yi I; Overall Christopher M; Stack M Sharon (a)

AUTHOR ADDRESS: (a)Dept. of Cell and Molecular Biology, Northwestern University Medical School, 303 E. Chicago Ave., Tarry 8-715, Chicago, IL, 60611: mss130@northwestern.edu**USA

JOURNAL: Journal of Biological Chemistry 276 (27):p24833-24842 July 6,

2001

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MEDIUM: print ISSN: 0021-9258

DOCUMENT TYPE: Article RECORD TYPE: Abstract LANGUAGE: English

SUMMARY LANGUAGE: English

ABSTRACT: Type I collagen stimulation of pro-matrix metalloproteinase (pro-MMP)-2 activation by ovarian *cancer* cells involves betal integrin receptor clustering; however, the specific cellular and biochemical events that accompany MMP processing are not well characterized. Collagenolysis is not required for stimulation of pro-MMP-2 activation, and denatured collagen does not elicit an MMP-2 activation response. Similarly, DOV13 cells bind to intact collagen utilizing both alpha2beta1 and alpha3beta1 integrins but interact poorly with collagenase-treated or thermally denatured collagen. *Antibody*-induced clustering of alpha3beta1 strongly promotes activation of pro-MMP-2, whereas alpha2beta1 integrin clustering has only marginal effects. Membrane-type 1 (MT1)-MMP is present on the DOV13 cell surface as both an active 55-kDa TIMP-2-binding species and a stable catalytically inactive 43-kDa form. Integrin clustering stimulates cell surface expression of MT1-MMP and co-localization of the proteinase to aggregated integrin complexes. Furthermore, cell surface proteolysis of the 55-kDa MT1-MMP species occurs in the absence of active MMP-2, suggesting MT1-MMP autolysis. Cellular invasion of type I collagen matrices requires collagenase activity, is blocked by tissue *inhibitor* of metalloproteinases-2 (TIMP-2) and collagenase-resistant collagen, is unaffected by TIMP-1, and is accompanied by pro-MMP-2 activation. Together, these data indicate that integrin stimulation of MT1-MMP activity is a rate-limiting step for type I collagen invasion and provide a mechanism by which this activity can be down-regulated following collagen clearance.

REGISTRY NUMBERS: 9001-12-1: COLLAGENASE; 146480-35-5: MATRIX
METALLOPROTEINASE-2; 146480-35-5: MMP-2; 161384-17-4: MEMBRANE-TYPE
1-MATRIX METALLOPROTEINASE; 161384-17-4: MT1-MMP; 148969-98-6:
PRO-MATRIX METALLOPROTEINASE-2; 148969-98-6: PRO-MMP-2; 140208-24-8:
TIMP-1; 124861-55-8: TIMP-2

DESCRIPTORS:

MAJOR CONCEPTS: Biochemistry and Molecular Biophysics; Metabolism
BIOSYSTEMATIC NAMES: Hominidae--Primates, Mammalia, Vertebrata, Chordata,

ORGANISMS: DOV13 cell line (Hominidae) -- ovarian *cancer* cells
BIOSYSTEMATIC CLASSIFICATION (SUPER TAXA): Animals; Chordates; Humans;
Mammals; Primates; Vertebrates

CHEMICALS & BIOCHEMICALS: alpha-2beta-1 integrin; alpha-3beta-1 integrin; collagenase; matrix metalloproteinase-2 {MMP-2}; membrane-type 1-matrix metalloproteinase {MT1-MMP}--cell surface expression, cell surface proteolysis; pro-matrix metalloproteinase-2 { pro-MMP-2}--activation; tissue *inhibitor* of metalloproteinases-1 { TIMP-1}; tissue *inhibitor* of metalloproteinases-2 {TIMP-2}; *type I

er 4) 2 44 collagen* MISCELLANEOUS TERMS: collagenolysis; functional interplay CONCEPT CODES: Cytology and Cytochemistry-Animal 02506 Cytology and Cytochemistry-Human 02508 Biochemical Studies-General 10060 Biochemical Studies-Proteins, Peptides and Amino Acids 10064 Enzymes-General and Comparative Studies; Coenzymes 10802 13002 Metabolism-General Metabolism; Metabolic Pathways BIOSYSTEMATIC CODES: 86215 Hominidae

22/5/3 (Item 3 from file: 5) DIALOG(R) File 5:Biosis Previews(R) (c) 2002 BIOSIS. All rts. reserv.

10838112 BIOSIS NO.: 199799459257

Transforming growth factor beta upregulates the integrin-mediated adhesion of human prostatic carcinoma cells to type I collagen.

AUTHOR: Kostenuik Paul J; Singh Gurmit; Orr F William(a) AUTHOR ADDRESS: (a) Dep. Pathol., Univ. Manitoba Health Sci. Cent., 820 Sherbrook St., Winnipeg, MB R3A 1R9**Canada JOURNAL: Clinical & Experimental Metastasis 15 (1):p41-52 1997 ISSN: 0262-0898

RECORD TYPE: Abstract LANGUAGE: English

ABSTRACT: Prostate *cancer* frequently metastasizes to bone, and we propose that this process may be facilitated by the adhesion of metastatic cells to bone-derived type I collagen. We examined collagen receptor function and regulation in osteotropic PC-3 human prostatic carcinoma cells. PC-3 cell adhesion to immobilized human type I collagen was promoted by Mn-2+ and Mg-2+ ions and was RGD-independent. *Antibodies* directed against beta-1 or alpha-2 integrin subunits *inhibited* adhesion to collagen by 90% and 53%, respectively, suggesting involvement of the alpha-2-beta-1 receptor. Anti-alpha-1 or anti-alpha-3 *antibodies* had no effect on adhesion. Flow cytometry and immunoprecipitation of (35S) methionine-labeled cells demonstrated that alpha-2-beta-1 was the major collagen receptor expressed by PC-3 cells. The pretreatment of PC-3 cells with transforming growth factor-beta-1 (TGF-beta-1), a major bone-derived growth factor, caused a rapid (2 h) 2-fold increase in the de novo synthesis of alpha-2 and beta-1 integrin subunits, and also increased by 2- to 3-fold the adhesion and spreading of PC-3 cells on collagen. We conclude that alpha-2-beta-1 is the major collagen receptor employed by PC-3 cells, and that alpha-2-beta-1 upregulation by TGF-beta is associated with an increased adhesion and spreading on collagen. The data suggest that exposure of metastatic PC-3 cells to the high levels of TGF-beta in bone may promote their ability to adhere to bone-derived collagen, which may thereby facilitate the localization of metastatic cells in the skeleton.

REGISTRY NUMBERS: 153-87-7Q: INTEGRIN; 60791-49-3Q: INTEGRIN DESCRIPTORS:

MAJOR CONCEPTS: Biochemistry and Molecular Biophysics; Cell Biology; Endocrine System (Chemical Coordination and Homeostasis); Membranes (Cell Biology); Oncology (Human Medicine, Medical Sciences); Reproductive System (Reproduction); Skeletal System (Movement and Support); Urology (Human Medicine, Medical Sciences) BIOSYSTEMATIC NAMES: Hominidae--Primates, Mammalia, Vertebrata, Chordata, ORGANISMS: PC-3 (Hominidae) -- cell line

BIOSYSTEMATIC CLASSIFICATION (SUPER TAXA): animals; chordates; humans; mammals; primates; vertebrates

CHEMICALS & BIOCHEMICALS: INTEGRIN

Research Article; BIOCHEMISTRY AND BIOPHYSICS; MISCELLANEOUS TERMS: BONE; BONE METASTASIS; HUMAN PROSTATIC CARCINOMA CELLS; INTEGRIN; INTEGRIN-MEDIATED ADHESION; NEOPLASTIC DISEASE; PROSTATE *CANCER*; PROSTATIC CARCINOMA CELLS; REPRODUCTIVE SYSTEM DISEASE/MALE; SKELETAL 1 3 ha SYSTEM; TRANSFORMING GROWTH FACTOR-BETA; TUMOR BIOLOGY; *TYPE I COLLAGEN*; UROLOGIC DISEASE CONCEPT CODES: Cytology and Cytochemistry-Human 02508 Biophysics-Molecular Properties and Macromolecules 10506 Biophysics-Membrane Phenomena 10508 Urinary System and External Secretions-Pathology 15506 Reproductive System-Pathology 16506 Endocrine System-General 17002 Bones, Joints, Fasciae, Connective and Adipose Tissue-Pathology 18006 Neoplasms and Neoplastic Agents-Pathology; Clinical Aspects; 24004 Systemic Effects Neoplasms and Neoplastic Agents-Biochemistry 24006 10064 Biochemical Studies-Proteins, Peptides and Amino Acids BIOSYSTEMATIC CODES: 86215 Hominidae

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Evidence for preferential adhesion of ovarian epithelial carcinoma cells to type I collagen mediated by the alpha-2-beta-1 integrin.

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ABSTRACT: Epithelial ovarian carcinoma, the leading cause of gynecologic *cancer* death, is characterized by widespread intra-abdominal metastases mediated primarily by surface shedding of tumor cells and peritoneal implantation. Whereas hematogenous metastasis is known to involve cellular adhesion, extracellular matrix proteolysis and cell migration, the role of these processes in the intraperitoneal dissemination of ovarian *cancer* remains unclear. To analyze further the role of adhesion and proteolysis in ovarian carcinoma dissemination, we have characterized the adhesive profiles of 4 primary cultures of ovarian carcinoma cells and 5 ovarian carcinoma cell lines. Our data demonstrate preferential adhesion of ovarian carcinoma cells to interstitial type I collagen. Analysis of adhesion molecule expression demonstrated the presence of the alpha-2 and beta-1 integrin subunits by cell surface ELISA, immunoprecipitation and immunohistochemistry. Furthermore, *antibodies* directed against the alpha-2 and beta-1 subunits *inhibited* adhesion of ovarian carcinoma cells to type I collagen by 56% and 95%, respectively. Plasminogen activator and matrix metalloproteinase production by adherent cells was not altered as a consequence of adhesion to individual extracellular matrix proteins; however, adhesion to an extracellular matrix comprised primarily of interstitial collagen increased plasminogen activator activity in 5 of 5 cell lines. Since the ovarian carcinoma micro-environment is rich in type I collagen, our data suggest that preferential adhesion to type I collagen followed by secretion of serine and metalloproteinases may represent a biochemical mechanism by which the intraperitoneal dissemination of ovarian carcinoma is mediated.

REGISTRY NUMBERS: 153-87-7Q: INTEGRIN; 60791-49-3Q: INTEGRIN; 9001-92-7: PROTEINASE DESCRIPTORS:

MAJOR CONCEPTS: Biochemistry and Molecular Biophysics; Cell Biology; Oncology (Human Medicine, Medical Sciences); Reproductive System (Reproduction)

BIOSYSTEMATIC NAMES: Hominidae--Primates, Mammalia, Vertebrata, Chordata, Animalia

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ORGANISMS: human (Hominidae)
 BIOSYSTEMATIC CLASSIFICATION (SUPER TAXA): animals; chordates; humans;
    mammals; primates; vertebrates
                             INTEGRIN; PROTEINASE
  CHEMICALS & BIOCHEMICALS:
 MISCELLANEOUS TERMS: ALPHA 2 BETA 1 INTEGRIN; DOV 13 CELL LINE; HUMAN
    OVARIAN CARCINOMA CELLS; HUMAN OVARIAN EPITHELIAL ASCITES CELLS;
    MATRIX-DEGRADING PROTEINASE; NEOPLASTIC DISEASE; OVARIAN CARCINOMA;
   OVARIAN CARCINOMA DISSEMINATION; OVARIAN METASTASES; OVCA 420 CELL LINE
    ; OVCA 429 CELL LINE; OVCA 432 CELL LINE; OVCA 433 CELL LINE;
    PERITONEAL; PRIMARY OVARIAN EPITHELIAL TUMOR; REPRODUCTIVE SYSTEM;
    REPRODUCTIVE SYSTEM DISEASE/FEMALE; SECRETION; TUMOR BIOLOGY; *TYPE I
    COLLAGEN*
CONCEPT CODES:
          Cytology and Cytochemistry-Human
  02508
          Biochemical Studies-Proteins, Peptides and Amino Acids
  10064
         Biochemical Studies-Carbohydrates
  10068
         Biophysics-Molecular Properties and Macromolecules
  10506
  16506
         Reproductive System-Pathology
         Neoplasms and Neoplastic Agents-Neoplastic Cell Lines
  24005
         Neoplasms and Neoplastic Agents-Biochemistry
 24006
BIOSYSTEMATIC CODES:
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Hominidae

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